### FIG. 1A

aagcggccggtctgcagtcggagacttgcaggcagcaaacacggtgcgaacgaa	90 180 232
M V V O T R F P S W I I L C Y I W L L G F A H T G E A O A A A A A A A A A A A A A A A A A	322
K E V L L D S K Å Q Q T E L E W I S Š P P S G W E E I S Ğ AAGGAAGTACTATTACTGGACTCGAAAGCACAACAAACAGAATTGGAATGGATTTCCTCTCCACCCAGTGGGTGG	412
L D E N Y T P I R T Y Q V C Q V M E P N Q N N W L R T N W : TTGGATGAGAACTACACTCCGATAAGAACATACCAGGTGTGCCAGGTCATGGAGCCCAACCAGAACAACTGGCTGCGGACTAACTGGATT	502
S K G N A Q R I F V E L K F T L R D © N S L P G V L G T © K TCTAAAGGCAACGCACAAAGGATTTTTGTAGAATTGAAATTCACCTTGAGGGATTGTAATAGTCTTCCCGGAGTCCTGGGAACTTGCAAG	592
130 150 E T F N L Y Y E T D Y D T G R N I R E N L Y V K I D T I A GAAACGTTTAATTTGTACTATTATGAAACAGACTACGACACCGGCAGGAATATACGAGAAAACCTTTATGTTAAAATAGACACCATTGCT	682
160 170 180  A D E S F T Q G D L G E R K M K L N T E V R E I G P L S K E GCAGATGAAAGTTTCACACAAGGTGACCTTGGTGAAAGAAGATGAAGCTGAACACTGAGGTGAGAGAGTTGGACCTTTGTCCAAAAAG	772
190 200 210 G F Y L A F Q D V G A C I A L V S V K V Y Y K K W T I V E GGATTCTATCTTGCCTTTCAGGATGTAGGGGCTTGCATAGCATTGGTTTCTGTCAAAGTGTACTACAAGAAGTGCTGGACCATTGTTGAG	862
N L A V F P D T V T G S E F S S L V E V R G T C V S S A E E AACTTAGCTGTCTTCCAGATACAGTGACTGGTTCGGAATTTTCCTCCTTAGTCGAGGTCCGTGGGACATGTGTCAGCAGTGCCGAGGAA	952
	1042
280 290 300 Q K G D T C E P C G R R F Y K S S S Q D L Q C S R C P T H S CAAAAAAGGGGACACTTGCGAACCCTGTGGCCGCAGGTTCTACAAATCTTCCTCTCAGGATCTCCAGTGTTCTCGTTGTCCAACCCACAGC	1132
	1222
340 350 360 PPSAPONLIFNIMOTITVSLEWSPPADNGGR CCTCCCTCTGCACCACAGAACCTTATTTTCAATATCAATCA	1312
370  N D V T Y R I L C K R C S W E Q G E C V P C G S N I G Y M P AACGATGTCACCTACAGAATACTGTGTAAGCGGTGCAGTTGGGAACAGGGAGAATGTGTGCCATGCGGAAGTAACATTGGATACATGCCC	1402
420 Q Q T G L E D N Y V T V M D L L A E A N Y T F E V E A V N G CAGCAGACGGGATTAGAGGATAACTATGTCACTGTCATGGACCTACTTGCCCATGCAAATTACACTTTCGAAGTTGAAGCTGTAAATGGA	1492
430 440 450 V S D L S R S Q R L F A A V S I T T G Q A A P S Q V S G V M GTTTCGGACTTAAGCAGATCCCAGAGGGCTCTTCGCTGCTGTTAGCATCACCACCGGTCAAGCAGCTCCCTCGCAAGTGAGTG	1582
460 470 480 K E R V L Q R S V Q L S W Q E P E E P N G V I T E Y E I K Y AAGGAGCGAGTACTGCAGCGGAGTGTGCAGCTTTCCTGGCAGGAGCCGGAGCATCCCAATGGAGTCATCACGGAATATGAAATCAAGTAT	1672
490 510 Y E K D Q R E R T Y S T L K T K S T S A S I N N L K P G T V TATGAGAAAGATCAACGGGAAAAGGACGTACTCAACACTCAAAACCAAGTCCACCTCCGCCTCCATTAATAATCTGAAACCGGGAACAGTG	1762
520 530 540 Y V F Q I R A V T A A G T G N Y S P R L D V A T L E E A S G TACGTCTTTCAGATCCGGGCGGTCACTGCCGGTTATGGAAACTACAGCCCTAGGCTTGATGTTGCCACACTTGAGGAAGCTTCAGGT	1852

## Title: METHOD FOR DIAGNOSIS AND TREATMENT OF MDK1 SIGNAL TRANSDUCTION DISORDERS

Inventor(s): Thomas CIOSSEK et al. Appl. No.: 08/368,776

FIG. 1B 550 560 570	
K M F E A T A V S S E O N P <u>V I I I A V V A V A G T I I L V</u> AAAATGTTTGAAGCGACAGCAGTCTCCAGTGAACAGAATCCTGTCATCATAATTGCTGTAGTGGCTGTAGCAGGGACCATCATCTTGGTG	1942
580 590 600  FMYFGFIIGRRHCGYSKADQEGDEELYFHF TTCATGGTGTTCGGCTCATCATTGGAAGAAGGCACTGTGGTTATAGCAAGGCTGACCAAGAAGGGGATGAAGAACTCTACTTTCATTTT	2030
610 620 630 KFPGTKTYIDPETYEDPNRAVHOFAKFIDA	-
AAATTTCCAGGCACCAAAACCTACATTGACCCTGAAACCTATGAGGACCCAAATAGAGCTGTCCATCAATTCGCCAAGGAGCTAGATGCC	2122
S C I K I E R V I G A G E F G E V C S G R L K L P G Q R D V TCCTGTATTAAAATTGAGCGTGTGATTGGTGCAGGAGAATTTGGAGAAGTTTGCAGTGGTCGTTTGAAACTTCCGGGCCAGAGAGATGTT	2212
A V A I K T L K V G Y T E K Q R R D F L C E A S I M G Q F D GCAGTGGCCATAAAAACCCTGAAAGTTGGTTACACAGAAAAGCAAAGGAGGGGACTTTTTATGCGAAGCAAGC	2302
700 710 720 H P N V V H L E G V V T R G K P V M I V I E F M E N G A L D CACCCAAATGTCGTCCATTTGGAAGGGGTTGTTACAAGAGGGAAGCCTGTCATGATTGTGATAGAGTTCATGGAGAATGGAGCCCTGGAT	2392
730 740 750  A F L R K H D G O F T V I Q L V G M L R G I A A G M R Y L A GCATTTCTCAGGAAACACGATGGGCAGTTTACAGTCATTCAGTTGGTAGGAATGTTGAGAGGGTATTGCCGCTGCCATGCGATACTTGGCT	2482
760 770 780  D M G Y V H R D L A A R N I L V N S N L V C K V S D F G L S GATATGGGATACGTTCACAGGGACCTTGCAGCGCGCAACATCCTTGTCAACAGCAATCTTGTTTGT	2572
790 800 810 R V I E D D P E A V Y T T T G G K I P V R W T A P E A I Q Y CGGGTTATAGAGGATGATCCCGAAGCTGTCTACACCACGACTGGTGGAAAAATTCCAGTAAGGTGGACTGCACCGGAAGCCATTCAATAC	2662
820 830 840 R K F T S A S D V W S Y G I V M W E V M S Y G E R P Y W D M CGGAAGTTCACCTCAGCCAGCGATGTGTGGGACTTGTCATGTGGGAAGTGATGTCTTATGGAGAAAGACCTTACTGGGACATG	2752
850 870 S N Q D V I K A I E E G Y R L P A P M D C P A G L H Q L M L TCAAATCAAGATGTCATTAAAGCGATAGAAGAAGGTTATCGTTTGCCGGCGCCCATGGATTGCCCAGCTGGTCTTCACCAGCTAATGCTG	2842
B80 900  D C W Q K D R A E R P K P E Q I V G I L D K M I R N R S S L GATTGTTGGCAGAAAGATCGGGCGGAAAGGCCAAAGTTTGAGCAGATAGTCGGAATTCTAGACAAAATGATTCGAAACCCAAGTAGTCTG	2932
910 920 930  K T P L G T C S R P L S P L L D Q S T P D F T A F C S V G E AAAACACCCCTGGGAACTTGTAGTAGACCCTTAAGCCCTCTTCTGGACCAGAGCACTCCTGACTTCACTGCCTTCTGTTCAGTTGGAGAA 940 950 950	3022
W L Q A I K M E R Y K D N F T A A G Y N S L E S V A R M T I TGGTTGCAAGCTATTAAAATGGAAAGGTATAAGGACAACTTCACAGCAGCGGGTTACAACTCACTC	3112
970 980 990 D D V M S L G I T L V G E Q K K I M S S I Q T M R A Q M L H GATGATGTGAGTTTAGGGATCACACTGGTTGGCCATCAAAAGAAGATCATGAGCAGCATCCAGACTATGCGGGCACAAATGTTGCAT	3202
L H G T G I O V * TTACACGGAACAGGCATCCAAGTGTGA	3229
cacatcggcctccctcagatgaggcttaagactgcaggagaacagttctggccttcagtatacgcatagaatgctgctagaagacagttgatatactgggtccttcct	3499
gttaaaatcaatgaaatattttccttaaaatatgtgatttcagactattcttttccagaaccatctgtgtttattctgcttaaggacttt gttttagaaagttatttgtagctttggaccttttagtgttaaatttatgacacgttactacactgggaacctttgaagactctcaaact taaaggaaagcaaaactacgcacatagtcgaggatggactttgtccttcatggctttggtatcctggctgtgtcattttgttaaaccagt gatgtttcatattgtttgctgattggcaggtagttcaaaattgcaagttgccaagagctctgatattttttaacaggatttttttt	3769 3859 3949 4039

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-11	
MDK1'.	

	2002	2092	2182	2272	2362	2452	2542	2632	2722	2812
290	S K A D 0 ragcaaggctgaccaa	actgtttaagacaat	ttattttgttgttat	naagacattgttctga	aattttcttcattgc	gaatttgatataaag	<b>ttgtttcaattttga</b>	ıtataaaccatttctc	atccagcctctatta	ıtactaattttatgta
580	V A V A G T I I L V F M V F G F I I G R R H C G Y S K A D Q GTGGCTGTAGCAGGGACCATCATCGTGTTCATGGTTTCGGCTTCATCGTAGGAAGGCACTGTGGTTATAGCAAGGCTGACCAA 600 610	E G D E E L Y F H S L V T N E H L S V L * GAAGGGGATGAAGAACTCTACTTC <u>IIIAGIAACAAATGAGCACCIGICAGTITTAIAA</u> accgcaacaataactgtttaagacaat	caattttggataaacaatcaactacagcagaataaatcaagatttttaagtcccattttcctttatacattctgcttatttgttgttatt	atgittaitititaaaacicigaicitgaitgaaigigataccaiaagcacagitaggcigcagigiaaaiaiataiaaagacaiigiiciga	gagcagtacgatttcatggaaagattgtttggtggctttgttaaaaattaataaagaatttltaaggatatagtgtaattttcttcattgc	attaatataaccaaatatgcctacctatctttgtcttgaaccaaatgaatagatttggaatactttattgtaattgaatttgatttgatataaag	ttgactgagcatttatgtgttacctgcatgcttctgggtgcattgaaatattttaacttttaaaatgatactatgttgtttcaattttga	ctaccttttgtgaggcatactggctacctcctctattagctaagatcttccaaagccttataatgaaaagtttatataaaccatttctc	tttcaaatcactgtcatacttggtcacggatcccaggaatattgtaaattttctaatttactctgcactttgtatatccagcctctatta	${\tt ccctcaaggtgaatataaaaactatgtcttttgaatatttctcttttgattttgttgatatagcagtccctcatatcttgtactaattttatgta}$
220	I L V F M V F G TCATCTTGGTGTTCGG 600	E G D E E L Y F H S L V T N E H L S V L * GAAGGGGATGAAGAACTCTACTTCATTCIIIAGIAACAAATGAGCACCTGICAGTTTTATA	aactacagcagaataaatcaagatt	tgatcttgattgaatgtgataccat	aaagattgtttggtggctttgttaa	cctacctatctttgtcttgaaccaa	ttacctgcatgcttctgggtgcatt	ctggctacctcctctattagctaa	ttggtcacggatcccaggaatattg	actatgtcttttgaatatttctctt
	V A V A G T I GTGGCTGTAGCAGGACCA	E G D E E L Y GAAGGGGATGAAGAACTCT,	caattttggataaacaatc	atgtttatttttaaactc	gagcagtacgatttcatgg	attaatataaccaaatatg	ttgactgagcatttatgtg	ctaccttttgtgaggcata	tttcaaatcactgtcatac	ccctcaaggtgaatataaa

2901

FIG. 2M

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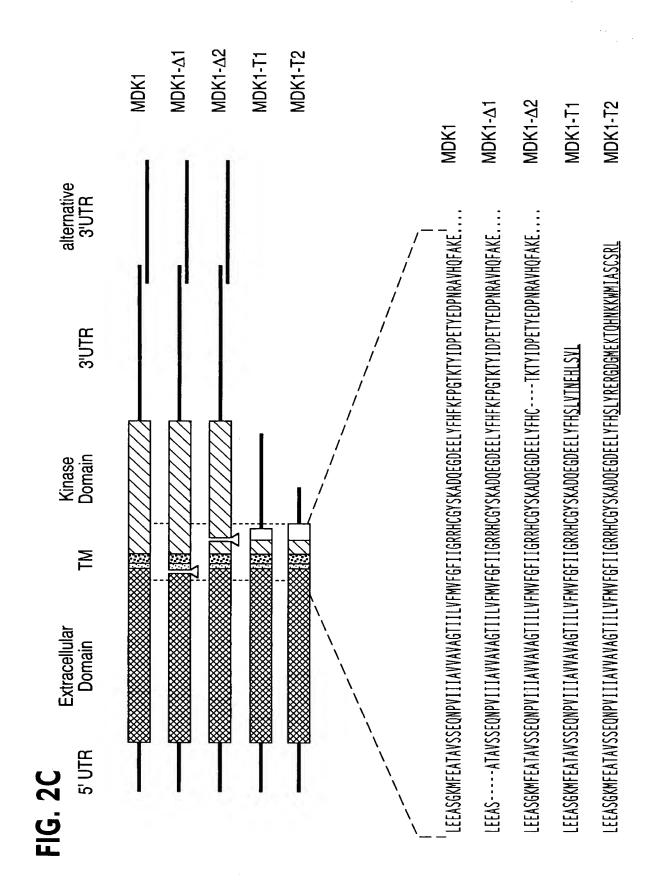
FIG. 28

MDK1-T2

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tttttgataattagtgctcagggggaggggggggaagtagagaaagcaaa Title: METHOD FOR DIAGNOSIS AND TREATMENT OF MDK1 SIGNAL TRANSDUCTION DISORDERS Inventor(s): Thomas CIOSSEK et al. Appl. No.: 08/368,776



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